

FIGURE 1A

ATGGGCGCACTGGCCGGCGCTGCTGCCTCTGCTGGCCAGTGGCTCCTGCGCGCC
M G A L A R A L L P L L A Q W L L R A
CCCCGGAGCTGGCCCCCGCGCCCTTCACGCTGCCCTCCGGGTGGCCCGGGCACGAAC
A P E L A P A P F T L P L R V A A A T N
CGCGTAGTTGCGCCCACCCGGGACCCGGGACCCCTGCCGAGCGCCACGCCACGGCTTG
R V V A P T P G P G T P A E R H A D G L
GCGCTGCCCTGGAGCCTGCCCTGGCGTCCCCCGCGGGCGCCGCAAACCTTGGCCATG
A L A L E P A L A S P A G A A A N F L A M
GTAGACAACCTGCAGGGGGACTCTGGCCGCGGCTACTACCTGGAGATGCTGATGGGACC
V D N L Q G D S G R G Y Y L E M L I G T
CCCCCGCAGAAGCTACAGATTCTCGTTGACACTGGAAGCAGTAACCTTGCCGTGGCAGGA
P P Q K L Q I L V D T G S S N F A V A G
ACCCCGCACT CCTACATAGACACGTACTTTGACACAGAGAGGTCTAGCACACATACCGCTCC
T P H S Y I D T Y F D T E R S S T Y R S
AAGGGCTTGACGTACAGTGAAGTACACACAAGGAAGCTGGACGGCTCGTTGGGAA
K G F D V T V K Y T Q G S W T G F V G E
GACCTCGTCACCATCCCCAAAGGCTCAATACTTCTTCTTGTCAACATTGCCACTATT
D L V T I P K G F N T S F L V N I A T I
TTTGAATCAGAGAATTCTTTGCTGGATTAAATGGAATGGAATACTTGGCCTAGCT
F E S E N F F L P G I K W N G I L G L A
TATGCCACACTTGCCAAGCCATCAAGTTCTGGAGACCTTCTCGACTCCCTGGTGACA
Y A T L A K P S S S L E T F F D S L V T
CAAGCAAACATCCCCAACGTTCTCCATGCAGATGTGTGGAGCCGGCTGCCCTTGCT
Q A N I P N V F S M Q M C G A G L P V A
GGATCTGGGACCAACGGAGGTAGTCTTGTCTGGGTGGAATTGAACCAAGTTGTATAAA
G S G T N G G S L V L G G I E P S L Y K
GGAGACATCTGGTATAACCCCTATTAAGGAAGAGTGGTACTACCAGATAGAAATTCTGAAA
G D I W Y T P I K E E W Y Y Q I E I L K
TTGGAAATTGGAGGCCAACGCTTAATCTGGACTGCAGAGAGTATAACGCCAGACAAGGCC
L E I G G Q S L N L D C R E Y N A D K A
ATCGTGGACAGTGGCACCAACGCTGCGCTGCCAGAAGGTGTTGATGCGGTGGTG
I V D S G T T L L R L P Q K V F D A V V
GAAGCTGTGGCCCCCGCATCTGATTCCAGAATTCTCTGATGGTTCTGGACTGGTCC
E A V A R A S L I P E F S D G F W T G S
CAGCTGGCGTGGACGAATTGGAAACACCTTGGTCTTACTTCCCTAAATCTCCATC
Q L A C W T N S E T P W S Y F P K I S I
TACCTGAGAGATGAGAACCTCAGCAGGTATTCCGTATCACAACTGCCAGCTTAC
Y L R D E N S S R S F R I T I L P Q L Y
ATTCAAGCCCATGATGGGGCCGGCTGAATTATGAATGTTACCGATTGGCATTCCCCA
I Q P M M G A G L N Y E C Y R F G I S P

FIGURE 1B

TCCACAAATGCGCTGGTATCGGTGCCACGGTGATGGAGGGCTTCTACGTCATCTTCGAC
S T N A L V I G A T V M E G F Y V I F D
AGAGCCCAGAAGAGGGTGGGCTTCGCAGCGAGCCCTGTGCAGAAATTGCAGGTGCTGCA
R A Q K R V G F A A S P C A E I A G A A
GTGTCTGAAATTCCGGCCTTCTAACAGAGGATGTAGCCAGCAACTGTGTCCCCGCT
V S E I S G P F S T E D V A S N C V P A
CAGTCTTGAGCGAGCCCATTGTGGATTGTGCCTATGCGCTCATGAGCGTCTGTGGA
Q S L S E P I L W I V S Y A L M S V C G
GCCATCCTCCTGTCTTAATCGTCCTGCTGCTGCCGTTCCGGTGTCAAGCGTCGCC
A I L L V L I V L L L P F R C Q R R P
CGTGACCTGAGGTCGTCAATGATGAGTCCTCTGGTCAGACATCGCTGGAAATGAATA
R D P E V V N D E S S L V R H R W K

GCCAGGCCTGACCTCAAGCAACCATGAACACTCAGCTATTAAGAAAATCACATTTCCAGGGC
AGCAGCCGGATCGATGGTGGCGCTTCTCCTGTGCCACCCGTCTCAATCTCTGTTCT
GCTCCCAGATGCCTTCTAGATTCACTGTCTTGATTCTTCAAGCTTCAAATC
CTCCCTACTCCAAGAAAAATAATTAAAAAAAAACTTCATTCTAAACCAAAAAAAA
AAAA

FIGURE 2A

ATGGCCCAAGCCCTGCCCTGGCTCCTGCTGTGGATGGCGCGGGAGTGCCTGCCAC
M A Q A L P W L L L W M G A G V L P A H
GGCACCCAGCACGGCATCCGGCTGCCCTGCGCAGCGGCCTGGGGGGCGCCCCCTGGGG
G T Q H G I R L P L R S G L G G G A P L G
CTGCGGCTGCCCGGGAGACCGACGAAGAGCCGAGGGAGCCGGCCGGAGGGCAGCTTT
L R L P R E T D E E P E E P G R R G S F
GTGGAGATGGTGGACAACCTGAGGGCAAGTCGGGGCAGGGCTACTACGTGGAGATGACC
V E M V D N L R G K S G Q G Y Y V E M T
GTGGCAGCCCCCGCAGACGCTAACATCCTGGTGGATAACAGGCAGCAGTAACTTGCA
V G S P P Q T L N I L V D T G S S N F A
GTGGGTGCTGCCCGGACCCCTTCATCGCTACTACCAAGGGCAGCTGTCCAGCACA
V G A A P H P F L H R Y Y Q R Q L S S T
TACCGGGACCTCCGAAGGGTGTGTATGTGCCCTACACCCAGGGCAAGTGGGAAGGGAG
Y R D L R K G V Y V P Y T Q G K W E G E
CTGGGCACCGACCTGGTAAGCATCCCCATGGCCCCAACGTCACTGTGCGTGCCAACATT
L G T D L V S I P H G P N V T V R A N I
GCTGCCATCACTGAATCAGACAAGTTCTCATCAACGGCTCCAACGGAAAGGCATCCTG
A A I T E S D K F F I N G S N W E G I L
GGGCTGGCTATGCTGAGATTGCCAGGCTTGTGGTGTGGCTTCCCTCAACCAGTCT
G L A Y A E I A R L C G A G F P L N Q S
GAAGTGCTGGCTCTGTCGGAGGGAGCATGATCATTGGAGGTATCGACCACCGCTGTAC
E V L A S V G G S M I I G G I D H S L Y
ACAGGCAGTCTGGTATACACCCATCCGGCGGGAGTGGTATTATGAGGTGATCATTGTG
T G S L W Y T P I R R E W Y Y E V I I V
CGGGTGGAGATCAATGGACAGGATCTGAAAATGGACTGCAAGGAGTACAACATGACAAG
R V E I N G Q D L K M D C K E Y N Y D K
AGCATTGGACAGTGGCACCAACCTCGTTGCCCAAGAAAGTGTGAAAGCTGCA
S I V D S G T T N L R L P K K V F E A A
GTCAAATCCATCAAGGCAGCCTCCACGGAGAAGTCCCTGATGGTTCTGGCTAGGA
V K S I K A A S S T E K F P D G F W L G
GAGCAGCTGGTGTGGCAAGCAGGCACCACCCCTGGAACATTTCAGTCATCTCA
E Q L V C W Q A G T T P W N I F P V I S
CTCTACCTAATGGGTGAGGTTACCAACCAGTCCTCCGCATCACCACCTCCGCAGCAA
L Y L M G E V T N Q S F R I T I L P Q Q
TACCTGGGCCAGTGGAAAGATGTGCCACGTCCCAGACGACTGTTACAAGTTGCCATC
Y L R P V E D V A T S Q D D C Y K F A I
TCACAGTCATCCACGGCACTGTTATGGGAGCTGTTATCATGGAGGGCTTCTACGTTGTC
S Q S S T G T V M G A V I M E G F Y V V
TTTGATGGGCCAAAACGAATTGGCTTGCTGTCAGCGCTGCCATGTGCACGATGAG
F D R A R K R I G F A V S A C H V H D E

FIGURE 2B

TTCAGGACGGCAGCGGTGGAAGGCCCTTGTACCTGGACATGGAAGACTGTGGCTAC
F R T A A V E G P F V T L D M E D C G Y
AACATTCCACAGACAGATGAGTCAACCTCATGACCATAGCCTATGTCATGGCTGCCATC
N I P Q T D E S T L M T I A Y V M A A I
TGCGCCCTTTCATGCTGCCACTCTGCCTCATGGTGTGTCAGTGGCGCTGCCTCCGCTGC
C A L F M L P L C L M V C Q W R C L R C
CTGCGCCAGCAGCATGATGACTTTGCTGATGACATCTCCCTGCTGAAGTGAGGAGGCCA
L R Q Q H D D F A D D I S L L K

TGGGCAGAAGATAGAGATTCCCTGGACCACACCTCCGTGGTCACTTGGTCACAAGTA
GGAGACACAGATGGCACCTGTGGCCAGAGCACCTCAGGACCCCTCCCCACCAAATGC
CTCTGCCTTGATGGAGAAGGAAAAGGCTGGCAAGGTGGGTTCCAGGGACTGTACCTGTAG
GAAACAGAAAAGAGAAGAAGAACGACTCTGCTGGCGGGAAATACTCTTGGTCACCTCAAA
TTAAGTCGGAAATTCTGCTGCTTGAAACTTCAGCCCTGAACCTTGTCCACCATTCC
TTAAATTCTCAACCAAAGTATTCTCTTTCTTAGTTCAGAAGTACTGGCATCACAC
GCAGGTTACCTTGGCGTGTGTCCTGTGGTACCTGGCAGAGAAGAGACCAAGCTGTT
CCCTGCTGGCAAAGTCAGTAGGAGAGGATGCACAGTTGCTATTTGCTTAGAGACAGG
GACTGTATAAACAAAGCCTAACATTGGTCAAAGATTGCCTTTGAAAAA

FIGURE 3A

ATGGCCCAAGCCCTGCCCTGGCTCTGCTGTGGATGGCGCGGGAGTGCCTGCCAC
M A Q A L P W L L L W M G A G V L P A H
GGCACCCAGCAGGCATCCGGCTGCCCTGCAGCGGCCTGGGGGCGCCCCCTGGGG
G T Q H G I R L P L R S G L G G A P L G
CTGCGGCTGCCCGGGAGACCGACGAAGAGCCCGAGGAGCCGGCCGGAGGGCAGCTT
L R L P R E T D E E P E E P G R R G S F
GTGGAGATGGTGGACAACCTGAGGGCAAGTCGGGGCAGGGCTACTACGTGGAGATGACC
V E M V D N L R G K S G Q G Y Y V E M T
GTGGGCAGCCCCCGCAGACGCTAACATCCTGGTGGATAACAGGCAGCAGTAACTTGCA
V G S P P Q T L N I L V D T G S S N F A
GTGGGTGCTGCCCTACCCCTCCTGCATCGCTACTACCAAGAGGCAGCTGTCCAGCACA
V G A A P H P F L H R Y Y Q R Q L S S T
TACCGGGACCTCCGGAAAGGGTGTGTATGTGCCCTACACCCAGGGCAAGTGGGAAGGGAG
Y R D L R K G V Y V P Y T Q G K W E G E
CTGGGCACCGACCTGGTAAGCATCCCCATGGCCCCAACGTCACTGTGCGTGCCAACATT
L G T D L V S I P H G P N V T V R A N I
GCTGCCATCACTGAATCAGACAAGTTCTCATCAACGGCTCCAACGGAAAGGCATCCTG
A A I T E S D K F F I N G S N W E G I L
GGGCTGGCCTATGCTGAGATTGCCAGGCCTGACGACTCCCTGGAGCCTTCTTGACTCT
G L A Y A E I A R P D D S L E P F F D S
CTGGTAAAGCAGACCCACGTTCCCAACCTCTCCCTGCAGCTTGTTGGTGCTGGCTTC
L V K Q T H V P N L F S L Q L C G A G F
CCCCTCAACCAGTCTGAAGTGTGGCTCTGTCGGAGGGAGCATGATCATTGGAGGTATC
P L N Q S E V L A S V G G S M I I G G I
GACCACTCGCTGTACACAGGCAGTCTCTGGTATAACACCCATCCGGCGGGAGTGGTATTAT
D H S L Y T G S L W Y T P I R R E W Y Y
GAGGTCACTATTGTGCGGGTGGAGATCAATGGACAGGATCTGAAAATGGACTGCAAGGAG
E V I I V R V E I N G Q D L K M D C K E
TACAACATATGACAAGAGCATTGTGGACAGTGGCACCACCAACCTCGTTGCCAAGAAA
Y N Y D K S I V D S G T T N L R L P K K
GTGTTTGAAGCTGCAGTCAAATCCATCAAGGCAGCCTCCACGGAGAAGTCCCTGAT
V F E A A V K S I K A A S S T E K F P D
GGTTTCTGGCTAGGAGAGCAGCTGGTGTGGCAAGCAGGCACCACCCCTGGAACATT
G F W L G E Q L V C W Q A G T T P W N I
TTCCCAAGTCATCTCACTCTACCTAAATGGGTGAGGTTACCAACCAGTCCTCCGCATCACC
F P V I S L Y L M G E V T N Q S F R I T
ATCCTTCCGCAGCAATACTGCAGGCCAGTGGAAAGATGTGGCACGTCCCAAGACGACTGT
I L P Q Q Y L R P V E D V A T S Q D D C
TACAAGTTGCCATCTCACAGTCATCCACGGGCACTGTTATGGGAGCTGTTATCATGGAG
Y K F A I S Q S S T G T V M G A V I M E

FIGURE 3B

GGCTTCTACGTTGCTTGATCGGCCCGAAAACGAATTGGCTTGCTGTCAAGCGCTTGC
G F Y V V F D R A R K R I G F A V S A C
CATGTGCACGATGAGTCAGGACGGCAGCGGTGGAAGGCCCTTTGTCACCTGGACATG
H V H D E F R T A A V E G P F V T L D M
GAAGACTGTGGCTACAACATTCCACAGACAGATGAGTCAACCCTCATGACCATA GCCTAT
E D C G Y N I P Q T D E S T L M T I A Y
GTCATGGCTGCCATCTGCCCTCTCATGCTGCCACTCTGCCCTCATGGTGTGTCAGTGG
V M A A I C A L F M L P L C L M V C Q W
CGCTGCCCTCGCTGCCAGCAGCATGATGACTTGCTGATGACATCTCCCTGCTG
R C L R C L R Q Q H D D F A D D I S L L
AAGTGAGGAGGCCATGGCAGAAGATAGAGATTCCCTGGACCACACCTCCGTGGTTCA
K

CTTTGGTCACAAGTAGGAGACACAGATGGCACCTGTGCCAGAGCACCTCAGGACCCCTCC
CCACCCACCAAATGCCCTGCCTGATGGAGAAGGAAAAGGCTGGCAAGGTGGTCCAG
GGACTGTACCTGTAGGAAACAGAAAAGAGAAGAAAGAACACTCTGCTGGCGGGAAATACT
CTTGGTCACCTCAAATTAAAGTCGGAAATTCTGCTGCTTGAAACTTCAGCCCTGAACCT
TTGTCACCATTCTTAAATTCTCCAACCAAAGTATTCTCTTTCTTAGTTCAGAA
GTACTGGCATCACACGCAGGTTACCTGGCGTGTGCCCTGTGGTACCTGGCAGAGAAG
AGACCAAGCTGTTCCCTGCTGCCAAAGTCAGTAGGAGAGGATGCACAGTTGCTATT
TGCTTAGAGACAGGGACTGTATAAACACAAGCCTAACATTGGTCAAAGATTGCCTTGA
ATTAAAAAAAAAAAAAA

FIGURE 4

ATGGCCCCAGCGCTGCACTGGCTCCTGCTATGGTGGCTCGGAATGCTGCCAG
M A P A L H W L L L W V G S G M L P A Q
GGAACCCATCTCGGCATCCGGCTGCCCTTCGCAGCGGCCTGGCAGGGCACCCCTGGC
G T H L G I R L P L R S G L A G P P L G
CTGAGGCTGCCCGGGAGACTGACGAGGAATCGGAGGAGCCTGGCCGGAGAGGCAGCTT
L R L P R E T D E E S E E P G R R G S F
GTGGAGATGGTGGACAACCTGAGGGAAAGTCCGCCAGGGCTACTATGTGGAGATGACC
V E M V D N L R G K S G Q G Y Y V E M T
GTAGGCAGCCCCCACAGACGCTAACATCCTGGTGGACACGGGAGTAGTAACTTGCA
V G S P P Q T L N I L V D T G S S N F A
GTGGGGCTGCCACACCCTTCCTGCATCGCTACTACCAAGAGGCAGCTGTCCAGCACA
V G A A P H P F L H R Y Y Q R Q L S S T
TATCGAGACCTCGAAAGGGTGTGTATGTGCCCTACACCCAGGGCAAGTGGGAGGGGAA
Y R D L R K G V Y V P Y T Q G K W E G E
CTGGGCACCGACCTGGTGAGCATCCCTCATGGCCCAACGTCAGTGTGCGTGCACATT
L G T D L V S I P H G P N V T V R A N I
GCTGCCATCACTGAATCGGACAAGTCTTCATCAATGGTCCAAGTGGGAGGGCATCCTA
A A I T E S D K F F I N G S N W E G I L
GGGCTGGCTATGCTGAGATTGCCAGGCCGACGACTCTTGGAGCCCTTCTTGACTCC
G L A Y A E I A R P D D S L E P F F D S
CTGGTGAAGCAGACCCACATTCCAACATCTTCCCTGCAGCTCTGTGGCGCTGGCTTC
L V K Q T H I P N I F S L Q L C G A G F
CCCCTCAACCAGACCGAGGCACTGGCCTCGGTGGAGGGAGCATGATCATTGGTGGTATC
P L N Q T E A L A S V G G S M I I G G I
GACCACTCGCTATAACACGGGAGTCTCTGGTACACACCCATCCGGGGAGTGGTATTAT
D H S L Y T G S L W Y T P I R R E W Y Y
GAAGTGATCATTGTACGTGTGGAAATCAATGGTCAAGATCTCAAGATGGACTGCAAGGAG
E V I I V R V E I N G Q D L K M D C K E
TACAAC TACGACAAGAGCATTGTGGACAGTGGGACCACCAACCTCGCTGCCAAGAAA
Y N Y D K S I V D S G T T N L R L P K K
GTATTTGAAGCTGCCGTCAAGTCATCAAGGCAGCCTCTGCACGGAGAAGTCCCGGAT
V F E A A V K S I K A A S S T E K F P D
GGCTTTGGCTAGGGGAGCAGCTGGTGCTGGCAAGCAGGCACGACCCCTTGGAACATT
G F W L G E Q L V C W Q A G T T P W N I
TTCCCACTTCATTTACCTCATGGTGAAGTCACCAATCAGTCCTCCGCATCACC
F P V I S L Y L M G E V T N Q S F R I T
ATCCTTCCTCAGCAATA CCTACGGCGGTGGAGGACGTGGCCACGTCCAAGACGACTGT
I L P Q Q Y L R P V E D V A T S Q D D C
TACAAGTCGCTGCTCACAGTCATCCACGGCACTGTTATGGGAGCCGTACATGGAA
Y K F A V S Q S S T G T V M G A V I M E
GGTTCTATGCGTCTCGATCGAGCCGAAAGCGAATTGGCTTGCTGTCAGCGCTTGC
G F Y V V F D R A R K R I G F A V S A C
CATGTGCACGATGAGTTCAAGACGGCGAGTGGAAAGGTCCGTTGTTACGGCAGACATG
H V H D E F R T A A V E G P F V T A D M
GAAGACTGTGGCTACAACATTCCCCAGACAGATGAGTCACACTATGACCATAGCCTAT
E D C G Y N I P Q T D E S T L M T I A Y
GTCATGGCGCCATCTCGCCCTCTCATGGTGCACACTCTGCCCTATGGTATGTCAGTGG
V M A A I C A L F M L P L C L M V C Q W
CGCTGCCTGCCTGCCACAGCACGATGACTTGTGATGACATCTCCCTGCTC
R C L R C L R H Q H D D F A D D I S L L
AAGTAAGGAGGCTCGTGGCAGATGATGGAGACGCCCTGGACCACATCTGGTGGTCC
K
CTTGTCACATGAGTTGGAGCTATGGATGGTACCTGTGGCAGAGCACCTCAGGACCC
CACCAACCTGCCAATGCTTCTGGCGTGACAGAACAGAGAAATCAGGCAAGCTGGATTACA
GGGCTTGACCTGTAGGACACAGGAGAGGAAGGAAGCAGCGTTCTGGTGGCAGGAATAT
CCTTAGGCACCACAAACTTGAGTTGGAAATTGCTGCTGAGCTCAGCCCTGACCC
CTGCCCAAGCATTAGAGTCACCAACCTAAAGTATTCTTATGTCCTTCCAGAAGTAC
TGGCGTCAACTCAGGCTACCCGGCATGTGTCCCTGGTACCCCTGGCAGAGAAAGGGCC
AATCTCATTCCCTGCTGGCAAAGTCAGCAGAAGGTGAAGTTGCCAGTTGCTTAG
TGATAGGGACTGCAGACTCAAGCCTACACTGGTACAAAGACTGCGTCTTGAGATAACAA
GAA

FIGURE 5

1 MAQALPWLLLWMGAGVLP AHLTQHGIRLPLRSGLGGAPLGLRLPRETDEE 50
|| || | | | . . . | | | | | | | | | | | | | | | | | | | | | |
1 MAPALHWLLLWVGSGMLPAQGTHLGIRLPLRSGLAGPPLGLRLPRETDEE 50

51 PEEPGRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFA 100
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
51 SEEPGRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFA 100

101 VGAAPHPFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGETGTDLVSIPH 150
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
101 VGAAPHPFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGETGTDLVSIPH 150

151 GPNVTVRANIAAITESDKFFINGSNWEGILGLAYAEIARPDDSLEPFFDS 200
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
151 GPNVTVRANIAAITESDKFFINGSNWEGILGLAYAEIARPDDSLEPFFDS 200

201 LVKQTHVPNLFSIQLCGAGFPLNQSEVLASVGGSMIIGGIDHSLYTGSLW 250
| | | | : | | | | | | | | | | | | | | | | | | | | | | | | | | | |
201 LVKQTHIPNIFSLQLCGAGFPLNQTEALASVGGSMIIGGIDHSLYTGSLW 250

251 YTPIRREWYYEVIIVRVEINGQDLKMDCKE NYDKSIVDSGTTNLRLPKK 300
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
251 YTPIRREWYYEVIIVRVEINGQDLKMDCKE NYDKSIVDSGTTNLRLPKK 300

301 VFEAAVKSIAASSTEKF PDGF WLGEQLVCWQAGTT PWNI FPVI SLYLMG 350
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
301 VFEAAVKSIAASSTEKF PDGF WLGEQLVCWQAGTT PWNI FPVI SLYLMG 350

351 EVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAISQSSTGTVMGAVIME 400
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
351 EVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAVSQSSTGTVMGAVIME 400

401 GFYVVFDRAKRIGFAVSACHVHDEFRTAAVEGPFTLDMEDCGYNIPQT 450
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
401 GFYVVFDRAKRIGFAVSACHVHDEFRTAAVEGPFTADMEDCGYNIPQT 450

451 DESTLMTIAYVMAAICALFMLPLCLMVCQWRCLRCLRQQHDDFADDISLL 500
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
451 DESTLMTIAYVMAAICALFMLPLCLMVCQWRCLRCLRHQHDDFADDISLL 500

501 K 501
|
501 K 501

FIGURE 6A

ATGGCTAGCATGACTGGTGGACAGCAAATGGTCGGATCCACCCAGCACGGCATCCGG
M A S M T G G Q Q M G R G S T Q H G I R
CTGCCCTGCGCAGCGGCTGGGGCGCCCCCTGGGGCTGGCTGCCCGGGAGACC
L P L R S G L G G A P L G L R L P R E T
GACGAAGAGCCCAGGGAGCCGGGGAGGGCAGCTTGAGATGGTGGACAACCTG
D E E P E E P G R R G S F V E M V D N L
AGGGCAAGTCGGGCAGGGCTACTACGTGGAGATGACCGTGGCAGCCCCCGAGACG
R G K S G Q G Y Y V E M T V G S P P Q T
CTCAACATCCTGGTGGATAAGGCAGCAGTAACCTTGCACTGGGTGCTGCCACCC
L N I L V D T G S S N F A V G A A P H P
TTCCTGCATCGCTACTACCAGAGGCAGCTGCCAGCACATACCGGACCTCCGAAAGGGC
F L H R Y Y Q R Q L S S T Y R D L R K G
GTGTATGTGCCCTACACCCAGGGCAAGTGGGAAGGGAGCTGGCACCGACCTGGTAAGC
V Y V P Y T Q G K W E G E L G T D L V S
ATCCCCATGGCCCAACGTCACTGTGCGTCCAACATTGCTGCCATCACTGAATCAGAC
I P H G P N V T V R A N I A A I T E S D
AAGTTCTTCATCAACGGCTCCAACGGGAAGGCATCCTGGGCTGGCTATGCTGAGATT
K F F I N G S N W E G I L G L A Y A E I
GCCAGGCCTGACGACTCCCTGGAGCCTTCTTGACTCTCTGGTAAAGCAGACCGTT
A R P D D S L E P F F D S L V K Q T H V
CCCAACCTCTTCTCCCTGCAGCTTGAGCTGGCTGGCTCCCTCAACCAGTCTGAAGTG
P N L F S L Q L C G A G F P L N Q S E V
CTGGCCTCTGTCGGAGGGAGCATGATCATTGGAGGTATCGACCAACTCGCTGTACACAGGC
L A S V G G S M I I G G I D H S L Y T G
AGTCTCTGGTATAACACCCATCCGGGGAGTGGTATTATGAGGTATCATTGTGGGTG
S L W Y T P I R R E W Y Y E V I I V R V
GAGATCAATGGACAGGATCTGAAAATGGACTGCAAGGAGTACAACATGACAAGAGCATT
E I N G Q D L K M D C K E Y N Y D K S I
GTGGACAGTGGCACCACCAACCTCGTTGCCAAGAAAGTGGTAAAGCTGCAGTCAAA
V D S G T T N L R L P K K V F E A A V K
TCCATCAAGGCAGCCCTCCACGGAGAAGTCCCTGATGGTTCTGGCTAGGAGAGCAG
S I K A A S S T E K F P D G F W L G E Q
CTGGTGTGCTGGCAAGCAGGCACCACCCCTGGAACATTTCCAGTCATCTCACTCTAC
L V C W Q A G T T P W N I F P V I S L Y
CTAATGGGTGAGGTTACCAACCAGTCCTCCGCATCACCATCCTCCGCAGCAATACCTG
L M G E V T N Q S F R I T I L P Q Q Y L
CGGCCAGTGGAAAGATGTGGCCACGTCCCAAGACGACTGTTACAAGTTGCCATCTCACAG
R P V E D V A T S Q D D C Y K F A I S Q
TCATCCACGGGCACTGTTATGGAGCTGTTATCATGGAGGGCTTACGTTGTCTTGAT
S S T G T V M G A V I M E G F Y V V V F D

Inventors: Gurney et al.
Title: "Alzheimer's Disease Secretase, APP Substrates
Therefor, and Uses Therefor"
Atty. Ref.: 29915/6280N3
Phone No.: (312) 474-6300

Fig. 6B
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FIGURE 6B

CGGGCCCCAAAACGAATTGGCTTGCTGTCAGCGCTGCCATGTGCACGATGAGTTCA
R A R K R I G F A V S A C H V H D E F R
ACGGCAGCGGTGGAAGGCCCTTGTACCTGGACATGGAAGACTGTGGCTACAACATT
T A A V E G P F V T L D M E D C G Y N I
CCACAGACAGATGAGTCATGA
P Q T D E S *

FIGURE 7A

ATGGCTAGCATGACTGGTGGACAGCAAATGGTCGCGGATCGATGACTATCTCTGACTCT
M A S M T G G Q Q M G R G S M T I S D S
CCGCGTGAACAGGACGGATCCACCCAGCACGGCATCCGGCTGCCCTGCGCAGCGGCCTG
P R E Q D G S T Q H G I R L P L R S G L
GGGGCGCCCCCTGGGCTGCGCTGCCCGGGAGACCGACGAAGAGCCCAGGGAGGCC
G G A P L G L R L P R E T D E E P E E P
GGCCGGAGGGCAGCTTGAGATGGTGGACAACCTGAGGGCAAGTCGGGCAGGGC
G R R G S F V E M V D N L R G K S G Q G
TACTACGTGGAGATGACCGTGGCAGCCCCCGCAGACGCTAACATCCTGGTGGATACA
Y Y V E M T V G S P P Q T L N I L V D T
GGCAGCAGTAACCTTGAGCTGGCTGCCCTACCCCTGCATCGCTACTACCAG
G S S N F A V G A A P H P F L H R Y Y Q
AGGCAGCTGTCCAGCACATACCGGACCTCCGGAAGGGCGTGTATGTGCCCTACACCCAG
R Q L S S T Y R D L R K G V Y V P Y T Q
GGCAAGTGGAAAGGGAGCTGGCACCGACCTGGTAAGCATCCCCATGGCCCCAACGTC
G K W E G E L G T D L V S I P H G P N V
ACTGTGCGTGCCAACATTGCTGCCATCACTGAATCAGACAAGTTCTTCATCAACGGCTCC
T V R A N I A A I T E S D K F F I N G S
AACTGGGAAGGCATCCTGGGCTGGCTATGCTGAGATTGCCAGGCCTGACGACTCCCTG
N W E G I L G L A Y A E I A R P D D S L
GAGCCTTCTTGACTCTGGTAAAGCAGACCCACGTTCCAACCTCTCCCTGCAG
E P F F D S L V K Q T H V P N L F S L Q
CTTGAGCTGGCTGGCTCCCTCAACCAGTCTGAAGTGCTGGCTCTGGAGGGAGC
L C G A G F P L N Q S E V L A S V G G S
ATGATCATTGGAGGTATCGACCACTCGCTGTACACAGGCAGTCTCTGGTATAACACCCATC
M I I G G I D H S L Y T G S L W Y T P I
CGGCGGGAGTGGTATTATGAGGTATCATTGTGCGGGTGGAGATCAATGGACAGGATCTG
R R E W Y Y E V I I V R V E I N G Q D L
AAAATGGACTGCAAGGAGTACAACATATGACAAGAGCATTGTGGACAGTGGCACCACCAAC
K M D C K E Y N Y D K S I V D S G T T N
CTTCGTTGCCAACAGAAAGTGGTGAAGCTGCAGTCACCATCAAGGCAGCCTCC
L R L P K K V F E A A V K S I K A A S S
ACGGAGAAGTCCCTGATGGTTCTGGCTAGGAGAGCAGCTGGTGTGGCAAGCAGGC
T E K F P D G F W L G E Q L V C W Q A G
ACCACCCCTGGAACATTTCAGTCATCTCACTCTACCTAACCTAATGGGTGAGGTTACCAAC
T T P W N I F P V I S L Y L M G E V T N
CAGTCCTCCGCATACCATCCTCCGCAGCAATACCTGCAGGCCAGTGGAAAGATGTGGCC
Q S F R I T I L P Q Q Y L R P V E D V A
ACGTCCCAAGACGACTGTTACAAGTTGCCATCTCACAGTCATCCACGGGACTGTTATG
T S Q D D C Y K F A I S Q S S T G T V M

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Title: "Alzheimer's Disease Secretase, APP Substrates
Therefor, and Uses Therefor"
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Fig. 7B
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FIGURE 7B

GGAGCTTTATCATGGAGGGCTTCTACGTTGTCTTGATCGGGCCCGAAAACGAATTGGC
G A V I M E G F Y V V F D R A R K R I G
TTGCTGTCAGCGCTTGCCATGTGCACGATGAGTTCAAGACGGCAGCGGTGGAAGGCCCT
F A V S A C H V H D E F R T A A V E G P
TTTGTACCTTGGACATGGAAGACTGTGGCTACAACATTCCACAGACAGATGAGTCATGA
F V T L D M E D C G Y N I P Q T D E S *

FIGURE 8A

ATGACTCAGCATGGTATTCTGTCTGCCACTGCGTAGCGGTCTGGGTGGTGCCTCCACTGGGT
M T Q H G I R L P L R S G L G G A P L G -
CTGCGTCTGCCCGGGAGACCGACGAAGAGCCGAGGGAGGCCGGCCGGAGGGCAGCTTT
L R L P R E T D E E P E E P G R R G S F -
GTGGAGATGGTGGACAACCTGAGGGCAAGTCGGGGCAGGGCTACTACGTGGAGATGACC
V E M V D N L R G K S G Q G Y Y V E M T -
GTGGGAGCCCCCGCAGACGCTAACATCTGGTGATAACAGGCAGCAGTAACCTTGCA
V G S P P Q T L N I L V D T G S S N F A -
GTGGGTGCTGCCCGCACCCCTTCATCGCTACTACCAAGGGCAGCTGTCCAGCACA
V G A A P H P F L H R Y Y Q R Q L S S T -
TACCGGGACCTCCCGAAGGGCGTGTATGTGCCCTACACCCAGGGCAAGTGGGAAGGGGAG
Y R D L R K G V Y V P Y T Q G K W E G E -
CTGGGCACCGACCTGGTAAGCATCCCCATGGCCCCAACGTCACTGTGGGTGCCAACATT
L G T D L V S I P H G P N V T V R A N I -
GCTGCCATCACTGAATCAGACAAGTTCTTCATCAACGGCTCCAACGGAAAGGCATCCTG
A A I T E S D K F F I N G S N W E G I L -
GGGCTGGCCTATGCTGAGATTGCCAGGCCTGACGACTCCCTGGAGCCTTCTTGACTCT
G L A Y A E I A R P D D S L E P F F D S
CTGGTAAAGCAGACCCACGTTCCAACCTCTCCCTGCAGCTTGTGGTGTGGCTTC
L V K Q T H V P N L F S L Q L C G A G F -
CCCCTCAACCAGTCTGAAGTGTGGCTCTGTCGGAGGGAGCATGATCATTGGAGGTATC
P L N Q S E V L A S V G G S M I I G G I -
GACCACTCGCTGTACACAGGCAGTCTCTGGTATAACACCCATCCGGCGGGAGTGGTATTAT
D H S L Y T G S L W Y T P I R R E W Y Y -
GAGGTCACTATTGTGCCGGTGGAGATCAATGGACAGGATCTGAAAATGGACTGCAAGGAG
E V I I V R V E I N G Q D L K M D C K E
TACAACATATGACAAGAGCATTGTGGACAGTGGCACCACCAACCTCGTTGCCAAGAAA
Y N Y D K S I V D S G T T N L R L P K K -
GTGTTGAAGCTGCAGTCAAATCCATCAAGGCAGCCTCCACGGAGAAGTCCCTGAT
V F E A A V K S I K A A S S T E K F P D -
GGTTCTGGCTAGGAGAGCAGCTGGTGTGGCAAGCAGGCACCACCCCTGGAACATT
G F W L G E Q L V C W Q A G T T P W N I -
TTCCCAGTCATCTCACTCTACCTAATGGGTGAGGTTACCAACCAGTCCTTCGCATCACC
F P V I S L Y L M G E V T N Q S F R I T -
ATCCTTCCGCAGCAATACCTGCAGCCAGTGGAAAGATGTGGCCACGTCCAAGACGACTGT
I L P Q Q Y L R P V E D V A T S Q D D C -

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Fig. 8B
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FIGURE 8B

TACAAGTTGCCATCTCACAGTCATCCACGGGCACTGTTATGGGAGCTGTTATCATGGAG
Y K F A I S Q S S T G T V M G A V I M E -
GGCTTCTACGTTGTCTTGATCGGGCCCGAAAACGAATTGGCTTGCTGTCAGCGCTTGC
G F Y V V F D R A R K R I G F A V S A C -
CATTAG
H *

FIGURE 9

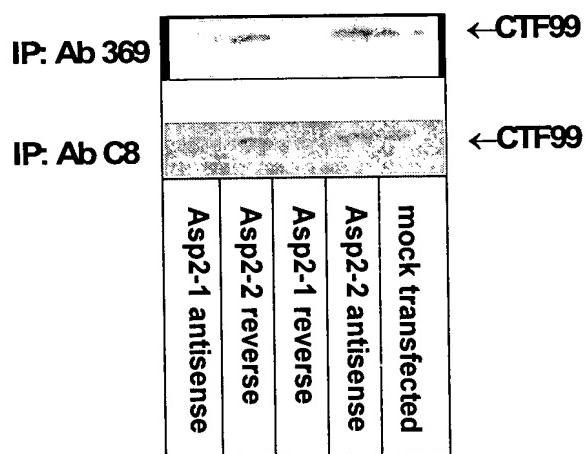
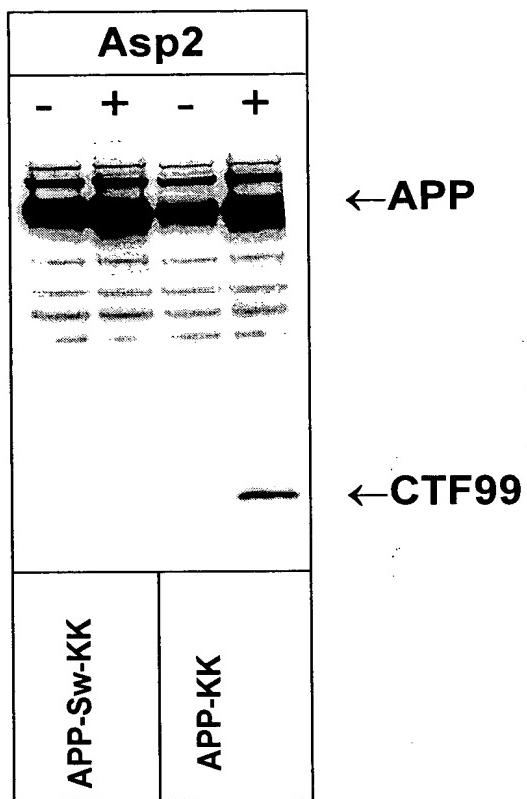


FIGURE 10



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Fig. 11
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FIGURE 11

MAQALPWLLLWMGAGVLPAG TQHGIRLPLRSGLGGAPLGLRLPRETDEE
PEEPGRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFA
VGAAPHFFLHRYYQRQLSSTYRDLRKGVYVPTYQGKWEGETGTDLVSIPH
GPNVTVRANIAAITESDKFFINGSNWEGLAYAEIARPDDSLPFFDS
LVKQTHVPNLFLQLCGAGFPLNQSEVLASVGGSMIIGGIDHSLYTGSILW
YTPIRREWYYEVIVRVEINGQDLKMDCKEENYDKSIVDSGTTNLRLPKK
VFEAAVKSIIKAASSTEKFPGFWLGEQLVCWQAGTPWNIFPVISLYLMG
EVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAISQSSTGTVMGAVIME
GFYVVVFDRARKRIGFAVSACHVDEFRTAAVEGPFTLDMEDCGYNIPQT
DES

FIGURE 12

MAQALPWLLLWMGAGVLPAG TQHGIRLPLRSGLGGAPLGLRLPRETDEE
PEEPGRGSFVEMVDNLRGKSGQGYYVEMTVGSPPQTLNILVDTGSSNFA
VGAAPHFLHRYYQRQLSSTYRDLRKGVYVPYTQGKWEGETGTDLVSIPH
GPNVTVRANIAAITESDKFFINGSNWEGLAYAEIARPDDSLPFFDS
LVKQTHVPNLFSLQLCGAGFPLNQSEVLASVGGSMIIGGIDHSLYTGSLW
YTPIRREWYYEVIVRVEINGQDLKMDCKEINYDKSIVDSGTTNLRLPKK
VFEAAVKSIAASSTEKFDPGFWLGEQLVCWQAGTTPWNIFPVISLYLMG
EVTNQSFRITILPQQYLRPVEDVATSQDDCYKFAISQSSTGTVMGAVIME
GFYVVVFDRARKRIGFAVSACHVDEFRTAAVEGPFTLDMEDCGYNIPQT
DESHHHHHH